PATENT COOPERATION TREATY

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INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

(Chapter II of the Patent Cooperation Treaty)

(PCT Article 36 and Rule 70)

Applicant's or agent's file reference B-15300 PCT FOR FURTI		R ACTION See Form PCT/IPEA/416							
International application No. PCT/IB2005/000506	International filing date 25.02.2005	(day/month/year)	Priority date (day/month/year) 26.02.2004						
International Patent Classification (IPC) or r	national classification and I	PC							
C08B37/00									
Applicant	1 500.								
PHARMA MEDICAL-LIMITED et al 92 8 15 CARDEROTO DE 701907									
1. This report is the international preliminary examination report, established by this International Preliminary Examining Authority under Article 35 and transmitted to the applicant according to Article 36.									
2. This REPORT consists of a total of 5 sheets, including this cover sheet.									
3. This report is also accompanied by ANNEXES, comprising:									
a. a. sent to the applicant and to the International Bureau) a total of sheets, as follows:									
sheets of the description, claims and/or drawings which have been amended and are the basis of this report and/or sheets containing rectifications authorized by this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions).									
sheets which supersede earlier sheets, but which this Authority considers contain an amendment that goes beyond the disclosure in the international application as filed, as indicated in item 4 of Box No. I and the									
Supplemental Box.	7		u of alastuania anuiou(a)\						
b. (sent to the International Bureau only) a total of (indicate type and number of electronic carrier(s)), containing a sequence listing and/or tables related thereto, in computer readable form only, as indicated in the Supplemental Box Relating to Sequence Listing (see Section 802 of the Administrative Instructions).									
4. This report contains indications relating to the following items:									
☐ Box No. I Basis of the opinion									
☐ Box No. II Priority									
☐ Box No. III Non-establishn	nent of opinion with rega	rd to novelty, inventive step and industrial applicability							
☐ Box No. IV Lack of unity of									
☐ Box No. V Reasoned state applicability; cit	Box No. V Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement								
☐ Box No. VI Certain docum	☐ Box No. VI Certain documents cited								
☐ Box No. VII Certain defects	☐ Box No. VII Certain defects in the international application								
☐ Box No. VIII Certain observations on the international application									
Date of submission of the demand		Date of completion of thi	s report						
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05.09.2005		18.01.2006							
Name and mailing address of the internatio	nal	Authorized Officer	nes Palon-						
preliminary examining authority: ———————————————————————————————————			and the state of t						
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INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

International application No. PCT/IB2005/000506

	Box No. I	Basis of the report
1.	With regard	d to the language , this report is based on the international application in the language in which it was so otherwise indicated under this item.
	☐ This rewhich	port is based on translations from the original language into the following language , is the language of a translation furnished for the purposes of:
	☐ pub	rnational search (under Rules 12.3 and 23.1(b)) dication of the international application (under Rule 12.4) drnational preliminary examination (under Rules 55.2 and/or 55.3)
2.	have been	It to the elements * of the international application, this report is based on (replacement sheets which furnished to the receiving Office in response to an invitation under Article 14 are referred to in this priginally filed" and are not annexed to this report):
	Description	, Pages
	1-18	as originally filed
	Claims, Nur	mbers
	1-36	as originally filed
	Drawings, S	Sheets
	1/7-7/7	as originally filed
	□ a sequ	ence listing and/or any related table(s) - see Supplemental Box Relating to Sequence Listing
3.	☐ the ☐ the ☐ the ☐ the	nendments have resulted in the cancellation of: description, pages claims, Nos. drawings, sheets/figs sequence listing (specify): table(s) related to sequence listing (specify):
4.	had not bee Supplemen the the the the the	port has been established as if (some of) the amendments annexed to this report and listed below en made, since they have been considered to go beyond the disclosure as filed, as indicated in the stal Box (Rule 70.2(c)). description, pages claims, Nos. drawings, sheets/figs sequence listing (specify): table(s) related to sequence listing (specify):
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International application No. PCT/IB2005/000506

Box No. V Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)

Yes: Claims

1-36

No: Claims

Inventive step (IS)

Yes: Claims

1-36

No: Claims

Industrial applicability (IA)

Yes: Claims

1-21

No: Claims 22-36

2. Citations and explanations (Rule 70.7):

see separate sheet

INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY (SEPARATE SHEET)

Re Item V

Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

Reference is made to the following documents:

- D1: CARNEY S L: "EFFECT OF DIACETYL RHEIN ON THE DEVELOPMENT OF EXPERIMENTAL OSTEOARTHRITIS A BIOCHEMICAL INVESTIGATION" December 1996, OSTEOARTHRITIS AND CARTILAGE, BAILLIERE TINDALL, LONDON, GB, PAGE(S) 251-261, XP001056066 ISSN: 1063-4584
- D2: BARBUCCI R ET AL: "Hyaluronic acid hydrogel in the treatment of osteoarthritis" December 2002, BIOMATERIALS, ELSEVIER SCIENCE PUBLISHERS BV., BARKING, GB, PAGE(S) 4503-4513, XP004377521 ISSN: 0142-9612

1. Novelty

D1 is concerned with the use of rhein in the treatment of joint disorders and D2 deals with the implementation of hyaluronic acid in the treatment of osteoarthritis.

None of these documents discloses the esterification of hyaluronic acid with rhein.

The subject-matter of claims 1-36 is novel over the cited state of the art (Article 33(2) PCT).

2. Inventive step

The hyaluronic acid esterified with rhein exhibits an improved pharmacological activity than hyaluronic acid or rhein alone (see page 17, line 14 - page 18, line 2, as well as figures 4 and 5).

The technical problem underlying the present application is the provision of a compound based on hyaluronic acid with improved pharmacological activity.

None of the cited documents cited alone or in combination would have prompted the

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skilled person seeking to solve the above-mentioned technical problem to esterify hyaluronic acid with rhein.

The subject-matter of claims 1-36 is therefore considered inventive (Article 33(3) PCT).

3. Industrial applicability

- **3.1.** The subject-matter of present claims 1-21 appears to comply with the requirements of industrial applicability as stipulated in Article 33(4) PCT.
- **3.2.** Claims 22-36 relate to subject-matter considered by this Authority to be covered by the provisions of Rule 67.1(iv) PCT. Consequently, no opinion will be formulated with respect to the industrial applicability of the subject-matter of these claims (Article 34(4)(a)(I) PCT).